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http://doegenomestolife.org/pubs.shtml

November 2005

# **DOE Office of Science: Continuing a Tradition of Achievement**

In 1986 the DOE Office of Science launched the Human Genome Project to understand, at the DNA level, the effects of energy production on human health. The benefits from this project far exceed the original goal. Today, DOE is poised to take the next vital steps to translate the genetic code in DNA into a new understanding of how life works and apply those biological processes to serve its challenging missions. DOE has the historic perspective, track record, and infrastructure to conduct the large-scale, complex, mission-driven science needed to achieve these goals. In the effort to understand biological systems, these assets and the Genomes to Life program will complement and extend the capabilities and efforts of the National Institutes of Health, the National Science Foundation, and other agencies and institutions around the world.

**Contacts for technical programmatic** information on GTL

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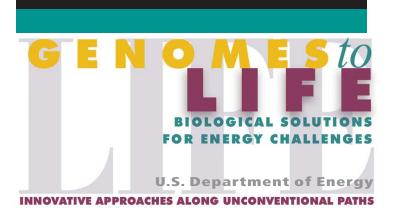
Copies of the April 2001 roadmap, along with associated documents, meeting reports, and image gallery, are downloadable via the Web:

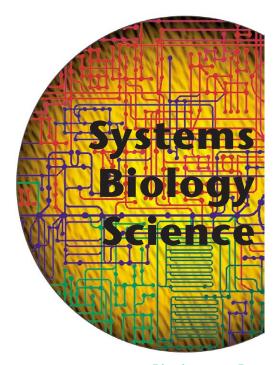
• DOEGenomesToLife.org

Future materials published about GTL may be requested from:

• Human Genome Management Information System

865/576-6669, Fax: /574-9888 mansfieldbk@ornl.gov

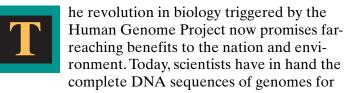




**Realizing the** Potential of the **Genome Revolution** 



# **Genomes to Life and Systems Biology Science**



many organisms—from microbes to plants to humans. For the first time, we can begin to explore the "operating systems" of life written into these genetic codes and put them to use. At the leading edge of this great scientific frontier is the Genomes to Life (GTL) program, whose overarching goal is to understand and use the diverse capabilities of microbes to target critical Department of Energy (DOE) mission challenges in energy security, global climate change, and toxic waste cleanup (see goals at right).

# A "Systems-Level" View

Although completed genomes provide "parts lists" for many microbes, we still have very little understanding of how these components create, sustain, and reproduce living systems. Obtaining this knowledge, a critical first step to harnessing microbial functions, requires extending the paradigm of comprehensive, whole-genome biology to the level of whole systems. In taking this approach, GTL uses DNA sequences as a departure point for explorations into the complex interactions of many levels of biological information, including genes, proteins, multimolecular assemblies ("molecular machines"), and the intricate labyrinth of pathways and networks in which they interact. These studies will enable an integrated and predictive understanding of how living cells function and respond to environmental changes. Revolutionary applications of this knowledge include clean, biology-based energy (e.g., methane or hydrogen), far more effective environmental cleanup with dramatic cost reduction, and reduced atmospheric carbon dioxide to mitigate global climate change.

# **Advanced Technology and Computing**

Just as the HGP required capabilities far beyond those available at its inception, concurrent technology development is needed to reach all GTL goals. Substantial efforts will be devoted, for example, to improving technologies for characterizing proteins and molecular machines, localizing them in cells and tissues,

carrying out high-throughput functional assays of complete cellular protein inventories, and sequencing and analyzing microbial DNA taken from natural environments.

Further, the wealth of data to be collected in GTL studies of dynamic living systems will be assimilated, understood, and modeled on the scale and complexity of real living systems and processes, an enormous task requiring the development of advanced computational methods and capabilities. These advances will complement experimental and theoretical biology to build a larger, intellectually richer, and more agile biology enterprise. These synergies are critical for making biological knowledge widely available and for stimulating new biological discovery and understanding.



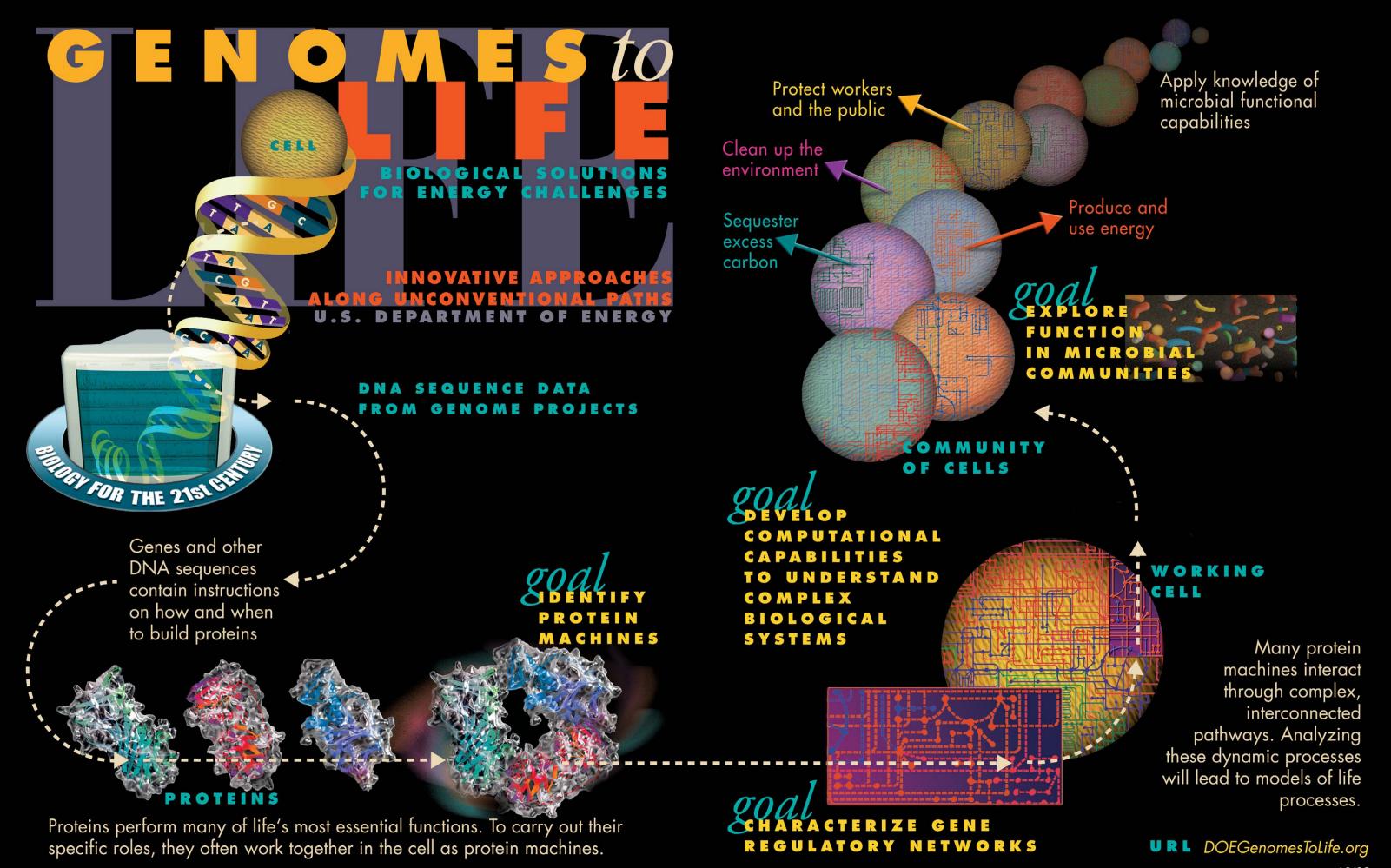
DNA sequence and high-throughput technologies goal 1 Identify and characterize the molecular machines of life goal 2 Characterize gene regulatory networks goal 3 Characterize the functional repertoire of complex microbial communities in their natural environments at the molecular level Develop the computational capabilities to advance understanding of complex biological systems and predict their behavior

**Systems** Biology **Understand how** molecular machines and other cell components function together in a living system

OBJECTIVES

- Contribute to a fundamental, comprehensive, and systematic understanding of life
- Support U.S. Department of Energy missions
- Environmental management
- Sustainable sources of energy
- Atmospheric and climate stability





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